AMENDMENT AND RESPONSE UNDER 37 CFR § 1.111

Serial Number: 09/730374 Filing Date: December 5, 2000

Title: USE OF GENETICALLY ENGINEERED ANTIBODIES TO CD38 TO TREAT MULTIPLE MYELOMA

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## **CLEAN VERSION OF PENDING CLAIMS**

FEB 2 0 2003

(Amended) A therapeutic composition, comprising:

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- a) a polypeptide comprising a polypeptide which specifically binds CD38 or a portion thereof linked to a polypeptide which specifically binds DNA or a portion thereof, wherein the polypeptide which specifically binds CD38 is an antibody; and
- b) a DNA sequence encoding a cytotoxic agent which is operably linked to a cell- or tissue-specific transcriptional unit.
- 3. (Amended) The composition of claim 1 wherein the antibody is obtained from an antibody secreted by hybridoma HB7.
- 4. The composition of claim 1 wherein the polypeptide which specifically binds DNA is protamine.
- 5. The composition of claim 1 wherein the cytotoxic agent is diphtheria toxin A chain, a cell suicide protein, Pseudomonas exotoxin, or an enzyme or protein that activates a chemotherapeutic agent.
- 6. The composition of claim 1 wherein the transcription unit is specific for B cells.
- 7. The composition of claim 1 wherein the transcription unit is specific for T cells.
- 8. The composition of claim 1 wherein the transcription unit is specific for myeloid cells.
- 9. (Amended) The composition of claim 1 wherein the antibody is a humanized antibody.
- 10. The composition of claim 1 further comprising a radioisotope linked to the fusion polypeptide.

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- 11. (Amended) The composition of claim 9 wherein the antibody is a scFv antibody.
- 12. (Amended) An isolated and purified polypeptide comprising at least a portion of a polypeptide that specifically binds CD38 and at least a portion of a polypeptide that specifically binds DNA, wherein the polypeptide which specifically binds CD38 is an antibody.
- 13. A method to inhibit the growth of CD38+ cells, comprising contacting cells in vitro with an effective amount of the composition of claim 1.
- 15. A method to inhibit or treat multiple myeloma, primary amyloidosis, monoclonal gammopathy, or acute myeloid leukemia, comprising: administering to a mammal in need of said treatment an effective amount of the composition of claim 1.
- 17. A recombinantly produced single chain fusion polypeptide comprising:
  - a) the Fv region of the light and the heavy chain of a CD38 specific antibody; and
- b) a DNA binding polypeptide, wherein the Fv region and the DNA binding polypeptide are recombinantly fused to form a single chain polypeptide that specifically binds CD38+ cells.
- 18. A pharmaceutical composition comprising a recombinantly produced single chain fusion polypeptide in a concentration sufficient to inhibit tumor cell growth, together with a pharmaceutically acceptable carrier wherein the fusion polypeptide comprises:
- a) a single chain Fv region of an antibody, wherein the Fv region comprises the VH and VL regions of the antibody; and
- b) a DNA binding polypeptide, wherein the Fv region and the DNA binding polypeptide are recombinantly fused to form a single molecule that specifically binds CD38+ cells.